IN THE CLAIMS:

1. (Currently Amended) A compound of formula:

$$(R^b)_p \\ N - L - B$$
 or
$$A = R^2$$

or a pharmaceutically acceptable salt, hydrate, <u>or solvate or prodrug</u> thereof, wherein

A represents a substituted or unsubstituted ring selected from the group consisting of an aromatic ring, a 5- or 6-membered heteroaromatic ring, a 5- or 6-membered cycloalkane ring and a 5- or 6-membered heterocycloalkane ring;

B is selected from the group consisting of cyclo(C_5 - C_8)alkyl, heterocyclo(C_5 - C_8)alkyl, cyclo(C_5 - C_8)alkenyl, heterocyclo(C_5 - C_8)alkenyl, aryl and heteroaryl;

L is (C₁-C₄)alkylene;

X and Y are <u>each</u> independently a divalent linkage selected from the group consisting of -O—; -C(O)—; $-N(R^3)$ —; $-C(O)N(R^3)$ —; $-S(O)_k$ —; $-SO_2N(R^3)$ —; and $-(C_1-C_2)$ alkylene—, wherein C_1 or C_2 is optionally substituted with $-OR^3$, $-N(R^3)COR^4$, $-C(O)NR^3R^4$, $-N(R^3)CO_2R^4$, $-N(R^3)C(O)N(R^4)R^5$, or -C(O); (C_1-C_2) alkylene, (C_1-C_2) alkylene— OR^3 , (C_1-C_2) alkylene— $N(R^3)COR^4$; (C_1-C_2) alkylene— $N(R^3)CO_2R^4$; (C_1-C_2) alkylene— $N(R^3)$;

 R^1 and R^2 are <u>each</u> independently selected from the group consisting of H, (C_1-C_4) alkyl, (C_2-C_8) alkenyl, (C_2-C_8) alkynyl, (C_1-C_8) heteroalkyl, aryl, aryl (C_1-C_4) alkyl, $-NR^6C(O)R^5$, $-C(O)R^5$ and $-NR^5C(O)NR^6$; $NR^6C(O)R^5$, $C(O)R^6$; and $NR^5C(O)NR^6$;

each R^b is selected from the group consisting of (C_1-C_4) alkyl, aryl, OR^7 , $C(O)R^7$ and $C(O)NR^7R^8$;

 R^3 and R^4 are independently selected from the group consisting of H, (C_1-C_8) alkyl, hetero (C_1-C_8) alkyl, aryl, aryl (C_1-C_4) alkyl, C(O)R', CO_2R' and C(O)NR'R'';

 R^5 , R^6 , R^7 and R^8 are independently selected from the group consisting of H, (C_1-C_8) alkyl, C(O)R''', CO_2R''' , aryl and aryl (C_1-C_4) alkyl;

optionally, R⁷ and R⁸ may be combined with the nitrogen to which each is attached to form a 5-, 6- or 7-membered ring;

R', R" and R" are independently selected from the group consisting of H, (C_1-C_8) alkyl, aryl and aryl (C_1-C_4) alkyl;

the subscript p is an integer of from 0 to 4; and the subscript k is an integer of from 0 to 2;

with the proviso that X and Y are not both $\underline{-O}$, $-N(R^3)$, $-S(O)_k$ or $SO_2N(R^3)$.

- 2. (Original) The compound of Claim 1, wherein the subscript p is 0.
- 3. (Withdrawn) The compound of Claim 1, wherein B is phenyl or naphthyl.
- 4. (Withdrawn) The compound of Claim 1, wherein B is selected from the group consisting of pyrrolyl, pyrazolyl, imidazolyl, pyrazinyl, oxazolyl, isoxazolyl, thiazolyl, furyl, thienyl, pyridyl, pyrimidyl, benzothiazolyl, purinyl, benzimidazolyl, indolyl, carbazolyl, indazolyl, carbolinyl, dibenzofuryl, dibenzothienyl, phenoxazinyl, phenothiazinyl, phenoxathiinyl, isoquinolyl, quinoxalinyl and quinolyl.
- 5. (Withdrawn) The compound of Claim 1, wherein B contains from 1 to 3 nitrogen atoms.
- 6. (Withdrawn) The compound of Claim 5, wherein B is selected from the group consisting of indolyl, carbazolyl and carbolinyl.
- 7. (Withdrawn) The compound of Claim 6, wherein B is 3-carbazolyl.
- 8. (Withdrawn) The compound of Claim 6, wherein B is 5-indolyl.
- 9. (Withdrawn) The compound of Claim 1, wherein **A** represents benzene, cyclohexane or cyclohexene.
- 10. (Withdrawn) The compound of Claim 1, wherein A represents benzene and B is 3-carbazolyl.

11. (Withdrawn) The compound of Claim 1, having a formula selected from the group consisting of:

12. (Withdrawn) The compound of Claim 1, having the formula (IV):

wherein:

each R^a is independently selected from the group consisting of halogen, halo(C_1 - C_4)alkyl, (C_1 - C_4)alkoxy, aryl(C_1 - C_4)alkyl, OC(O) R^{17} , NR¹⁷ R^{18} , SR¹⁷, cyano, nitro, CO_2R^{17} , $CONR^{17}R^{18}$, C(O) R^{17} , OC(O)NR¹⁷ R^{18} , NR¹⁸C(O)R¹⁷, NR¹⁸CO₂R¹⁷, NR¹⁹C(O)NR¹⁷R¹⁸, S(O)_kR¹⁷, S(O)_kNR¹⁷R¹⁸, N₃, (C₄-C₈)cycloalkyl, (C₅-C₈)cycloalkenyl, aryl and heteroaryl;

R¹⁷, R¹⁸ and R¹⁹ are independently selected from the group consisting of H,

 (C_1-C_8) alkyl, (C_1-C_8) heteroalkyl, aryl (C_1-C_4) alkyl and aryl; and the subscript m is an integer of from 0 to 4.

- 13. (Withdrawn) The compound of Claim 12, wherein X or Y is (C₁-C₂)alkylene—OH.
- 14. (Withdrawn) The compound of Claim 12, wherein Y is CH—OH.
- 15. (Withdrawn) The compound of Claim 12, wherein X is (C_1-C_2) alkylene— $N(R^3)COR^4$.
- 16. (Withdrawn) The compound of Claim 12, wherein X is CH-N(R³)COR⁴ and Y is CH-OH.
- 17. (Withdrawn) The compound of Claim 12, wherein X is N(R³) and Y is C(O).
- 18. (Withdrawn) The compound of Claim 12, wherein X is (C_1-C_2) alkylene, $N(R^3)$, $C(O)N(R^3)$ or $S(O)_k$ and Y is (C_1-C_2) alkylene.
- 19. (Withdrawn) The compound of Claim 12, wherein X is (C_1-C_2) alkylene and Y is C(O), $N(R^3)$, $C(O)N(R^3)$ or $S(O)_k$.
- 20. (Original) The compound of Claim 1 having the formula (V):

$$(R^a)_m$$
 R^2 R^2

 \mathbf{V}

- 21. (Original) The compound of Claim 20, wherein R^1 and R^2 are H.
- 22. (Original) The compound of Claim 1, having the formula:

VII

VIII

wherein

 R^{11} is selected from the group consisting of H, (C_1-C_4) alkyl, (C_2-C_8) alkenyl, (C_2-C_8) alkynyl, (C_1-C_8) heteroalkyl, aryl, aryl (C_1-C_4) alkyl, heteroaryl,

heteroaryl(C_1 - C_4)alkyl, (C_3 - C_8)cycloalkyl, (C_5 - C_8)cycloalkenyl, (C_3 - C_8)cycloalkyl-alkyl, (C_3 - C_8)cycloheteroalkyl, (C_3 - C_8)cycloheteroalkyl-alkyl, (C_3 - C_8)cycloheteroalkyl-alkyl

each R^c is independently selected from the group consisting of $(C_1\text{-}C_8)$ alkyl, $(C_2\text{-}C_8)$ alkenyl, $(C_2\text{-}C_8)$ alkynyl, $(C_1\text{-}C_8)$ heteroalkyl, halo $(C_1\text{-}C_8)$ alkyl, halogen, CN, NO_2 , OR^{14} , SR^{14} , $NR^{14}R^{15}$, $(C_3\text{-}C_8)$ cycloalkyl, $(C_5\text{-}C_8)$ cycloalkenyl, $(C_3\text{-}C_8)$ cycloalkyl-alkyl, $(C_3\text{-}C_8)$ cycloheteroalkyl-alkyl, $C(O)R^{14}$, CO_2R^{14} , $C(O)NR^{14}R^{15}$, aryl, aryl $(C_1\text{-}C_4)$ alkyl, heteroaryl, heteroaryl $(C_1\text{-}C_4)$ alkyl, $S(O)_kR^{14}$, $S(O)_kNR^{14}R^{15}$, $N(R^{15})S(O)_kR^{14}$, $OC(O)R^{14}$, OCO_2R^{14} , $OC(O)NR^{14}R^{15}$, $N(R^{16})C(O)NR^{14}R^{15}$, $N(R^{15})C(O)R^{14}$ and $N(R^{15})CO_2R^{14}$;

optionally, any two adjacent R^c groups may be combined to form a fused aryl ring or (C_5-C_8) cycloalkyl ring;

 R^{12} , R^{13} , R^{14} , R^{15} and R^{16} are independently selected from the group consisting of H, (C_1-C_8) alkyl, (C_1-C_8) heteroalkyl, aryl (C_1-C_4) alkyl and aryl;

the subscript q is an integer of from 0 to 7; and the subscript k is an integer of from 1 to 2.

23. (Original) The compound of Claim 22, having the formula:

$$(R^{a})_{m} \xrightarrow{V-X} X X XI$$

wherein

T02-009-1/US

each R^a is independently selected from the group consisting of halogen, halo(C_1 - C_4)alkyl, (C_1 - C_4)alkoxy, aryl(C_1 - C_4)alkyl, OC(O) R^{17} , NR¹⁷ R^{18} , SR¹⁷, cyano, nitro, CO_2R^{17} , CONR¹⁷ R^{18} , C(O) R^{17} , OC(O)NR¹⁷ R^{18} , NR¹⁸C(O)R¹⁷, NR¹⁸CO₂R¹⁷, NR¹⁹C(O)NR¹⁷ R^{18} , S(O)_kNR¹⁷ R^{18} , N3, (C₄-C₈)cycloalkyl, (C₅-C₈)cycloalkenyl, aryl and heteroaryl;

 R^{17} , R^{18} and R^{19} are independently selected from the group consisting of H, (C₁-C₈)alkyl, (C₁-C₈)heteroalkyl, aryl(C₁-C₄)alkyl and aryl;

- 6 -

the subscript m is an integer of from 0 to 4; and each subscript k is an integer of from 1 to 2.

24. (Currently Amended) The compound of any one of Claims 1, [12,] 20 and 23, wherein L is methylene.

25. (Currently Amended) The compound of Claim 23, having the formula (Xa):

Xa

wherein

L is methylene; and

X and Y are independently selected from $\underline{-(C_1-C_2)}$ alkylene—, wherein $\underline{C_1}$ or $\underline{C_2}$ is optionally substituted with $\underline{-OR^3}$, $\underline{-N(R^3)COR^4}$, $\underline{-C(O)NR^3R^4}$ or $\underline{-N(R^3)C(O)N(R^4)R^5}$. (C_1-C_2) alkylene— (C_1-C_2)

(Original) The compound of Claim 25, having a formula selected from the group 26. consisting of:

•

27. (Original) A compound of formula:

or a pharmaceutically acceptable salt, hydrate, or solvate solvate or prodrug thereof, wherein R^{20} and R^{23} independently represent H or OR^3 ;

 R^{21} and R^{22} independently represent H, OR³, N(R³)COR⁴, C(O)NR³R⁴, N(R³)CO₂R⁴, N(R³)C(O)N(R⁴)R⁵, N(R³)R⁴, C(O)N(R³)R⁴, N(R³)C(O)R⁴, (CH₂)C(O)N(R³)(R⁴), (CH₂)CO₂R³, or (C₁-C₄)alkyl;

 $R^{11} \ represents \ H, (C_1-C_4) alkyl, (C_2-C_8) alkenyl, (C_2-C_8) alkynyl, (C_1-C_8) heteroalkyl, aryl, aryl(C_1-C_4) alkyl, heteroaryl, heteroaryl(C_1-C_4) alkyl, (C_3-C_8) cycloalkyl, \\$

 (C_5-C_8) cycloalkenyl, (C_3-C_8) cycloalkyl-alkyl, (C_3-C_8) cycloheteroalkyl,

 (C_3-C_8) cycloheteroalkyl-alkyl, $C(O)R^{12}$, CO_2R^{12} , $C(O)NR^{12}R^{13}$, $S(O)_kR^{12}$ or $S(O)_kNR^{12}R^{13}$;

 R^{12} and R^{13} independently represent H, (C_1-C_8) alkyl, (C_1-C_8) heteroalkyl, aryl (C_1-C_4) alkyl or aryl;

 R^3 and R^4 independently represent H, (C_1-C_8) alkyl, hetero (C_1-C_8) alkyl, aryl, aryl (C_1-C_4) alkyl, C(O)R', CO_2R' or C(O)NR'R''; and

R', R" and R"' are independently selected from the group consisting of H, (C_1-C_8) alkyl, aryl and aryl (C_1-C_4) alkyl.

- 28. (Original) The compound of Claim 27, wherein R²⁰ and R²³ each represent H, R²² represents OH, and R²¹ represents N(R³)C(O)R⁴.
- 29. (Original) The compound of Claim 27, wherein R²⁰ represents OH, and R²² and R²³ each represent H, and R²¹ represents C₂ alkyl.
- 30. (Original) The compound of Claim 27, wherein R^{20} , R^{22} , and R^{23} each represent H and R^{21} represents $N(R^3)C(O)R^4$.
- 31. (Original) The compound of Claim 27, wherein R^{20} , R^{22} , and R^{23} each represent H and R^{21} represents $(CH_2)CO_2R^3$.
- 32. (Original) The compound of Claim 27, wherein R²⁰, R²², and R²³ each represent H and R²¹ represents (CH₂)C(O)N(R³)(R⁴).
- 33. (Original) The compound of Claim 27, having a formula that is selected from the group consisting of:

- 34. (Cancelled)
- 35. (Cancelled)
- 36. (Cancelled)
- 37. (Cancelled)
- 38. (Cancelled)
- 39. (Cancelled)
- 40. (Cancelled)
- 41. (Cancelled)
- 42. (Currently Amended) The compound of any one of Claims Claim 25, 38 and 40, wherein L is methylene.

- 43. (Withdrawn) A pharmaceutical composition comprising a pharmaceutically acceptable carrier or excipient and a compound of Claim 1.
- (Currently Amended) A method of treating a metabolic disorder, condition or 44. disorder selected from the group consisting of obesity, diabetes, anorexia nervosa, bulimia, pain, cancers, asthma, Parkinson's disease, acute heart failure, congestive heart failure, hypotension, hypertension, urinary retention, osteoporosis, angina pectoris, myocardial infarction, stroke, ulcers, allergies, benign prostatic hypertrophy, migraine, vomiting, anxiety, schizophrenia, manic depression, depression, delirium, dementia, severe mental retardation, Huntington's disease, Gilles de la Tourette's Syndrome, Syndrome X, insulin resistance, hyperglycemia, hyperuricemia, hyperinsulinemia, hypercholesterolemia, hyperlipidemia, dyslipidemia, mixed dyslipidemia, hypertriglyceridemia, male sexual dysfunction, female sexual dysfunction, fever, inflammation, rheumatoid arthritis, atherosclerosis, Alzheimer's disease, epilepsy, autism, bipolar disorder, neuroses, substance abuse, generalized anxiety disorder, panic disorder, obsessive compulsive disorder, posttraumatic stress syndrome, gall-bladder disease, sleep apnea syndrome, narcolepsy, insomnia, Shy-Drager Syndrome, multiple sclerosis, dystonia, coronary artery disease, cardiomyopathy, cachexia, osteoarthritis, Prader Willi Syndrome, hypothyroidism, hypogonadism, hyperprolactinemia, traumatic brain injury, ischemic reperfusion iniury, aneurysm, spinal cord injury and Pickwick Syndrome, comprising administering to a subject in need thereof a therapeutically effective amount of a compound of Claim 1.
- 45. (Withdrawn) A method of treating a condition or disorder responsive to MCHR2 modulation, comprising administering to a subject in need thereof a therapeutically effective amount of a compound of Claim 1.
- 46. (Withdrawn) A method of treating an MCHR2-mediated condition or disorder, comprising administering to a subject in need thereof a therapeutically effective amount of a compound of Claim 1.
- 47. (Withdrawn) The method of Claim 45 or 46, wherein said condition or disorder is selected from the group consisting of obesity, diabetes, anorexia nervosa, bulimia, pain, cancers, asthma, Parkinson's disease, acute heart failure, congestive heart failure, hypotension, hypertension, urinary retention, osteoporosis, angina pectoris,

myocardial infarction, stroke, ulcers, allergies, benign prostatic hypertrophy, migraine, vomiting, anxiety, schizophrenia, manic depression, depression, delirium, dementia, severe mental retardation, Huntington's disease, Gilles de la Tourette's Syndrome, Syndrome X, insulin resistance, hyperglycemia, hyperuricemia, hyperinsulinemia, hypercholesterolemia, hyperlipidemia, dyslipidemia, mixed dyslipidemia, hypertriglyceridemia, male sexual dysfunction, female sexual dysfunction, fever, inflammation, rheumatoid arthritis, atherosclerosis, Alzheimer's disease, epilepsy, autism, bipolar disorder, neuroses, substance abuse, generalized anxiety disorder, panic disorder, obsessive-compulsive disorder, posttraumatic stress syndrome, gall bladder disease, sleep apnea syndrome, narcolepsy, insomnia, Shy-Drager Syndrome, multiple sclerosis, dystonia, coronary artery disease, cardiomyopathy, cachexia, osteoarthritis, Prader-Willi Syndrome, hypothyroidism, hypogonadism, hyperprolactinemia, traumatic brain injury, ischemic reperfusion injury, aneurysm, spinal cord injury and Pickwick Syndrome.

- 48. (Withdrawn) The method of any one of Claims 44, 45 and 46, wherein said compound is administered in combination with a second therapeutic agent.
- 49. (Withdrawn) The method of Claim 48, wherein said second therapeutic agent is selected from the group consisting of an anti-obesity agent, an anti-diabetic agent, a non-steroidal antiinflammatory agent, an opioid analgesic, an antineoplastic agent, a cholesterol lowering agent, an antithrombotic agent, an anticonvulsant, an antiphychotic agent, a cholinesterase inhibitor, an anticholinergic agent, a dopaminergic agent, interferon β, a multiple sclerosis therapeutic agent, an anti-anxiety agent, an antidepressant, a phophodiester V inhibitor, an α-2 adrenergic receptor antagonist and an MCHR1 antagonist.
- 50. (Withdrawn) A method of modifying feeding behavior, comprising administering to a subject an amount of a compound of Claim 1 effective to reduce or enhance food intake by the subject by at least 5%.
- 51. (Withdrawn) A method of reducing body mass, comprising administering to a subject an amount of a compound of Claim 1 effective to decrease the body mass of the subject by at least 5% of baseline.
- 52. (Withdrawn) The method of Claim 51, wherein the body mass of the subject is decreased by at least 10% of baseline.

- (Withdrawn) A method of modulating MCHR2 in a cell, comprising contacting a cell 53. with a compound of Claim 1.
- 54. (Withdrawn) A method for modulating MCHR2, comprising contacting a protein with a compound of Claim 1.
- (Withdrawn) The method of Claim 54, wherein said compound is an MCHR2 55. antagonist.
- 56. (Withdrawn) A method for identifying a compound that modulates signal transduction, comprising
 - contacting an isolated or recombinant MCHR2 polypeptide with a compound a)of Claim 1 under conditions suitable for MCHR2-mediated signal transduction;
 - measuring intracellular Ca²⁺, cAMP or IP₃ in the absence and presence of said b)compound; and
 - comparing intracellular Ca²⁺, cAMP or IP₃ levels in the absence and presence c)of said compound;

wherein an increase or a decrease in intracellular Ca²⁺, cAMP or IP₃ level in the presence of said compound indicates that said compound modulates signal transduction.

- 57. (Withdrawn) A method for identifying a compound that modulates signal transduction, comprising
 - contacting an isolated or recombinant MCHR2 polypeptide with an MCHR2 a)ligand in the absence and presence of a compound of Claim 1 under conditions suitable for G-protein coupling to said polypeptide;
 - b)measuring G-protein activation in the absence and presence of said compound; and
 - comparing G-protein activation in the absence and presence of said c)compound;

wherein an increase or a decrease in G-protein activation in the presence of said

compound indicates that said compound modulates signal transduction.

- 58. (Withdrawn) A method for identifying a compound that modulates MCHR2, comprising
 - a)- contacting an isolated or recombinant MCHR2 polypeptide with an MCHR2 ligand in the absence and presence of a compound of Claim 1 under conditions suitable for ligand binding to said polypeptide;
 - b)- measuring ligand binding to said polypeptide in the absence and presence of said compound; and
 - c)- comparing ligand binding to said polypeptide in the absence and presence of said compound;

wherein an increase or a decrease in ligand binding in the presence of said compound indicates that said compound modulates MCHR2.

- 59. (Withdrawn) A method for identifying a compound that modulates MCHR2, comprising
 - a)- contacting a cell comprising a target gene that is activated by an MCHR2 ligand with a compound of Claim 1 under conditions suitable for transcription or expression of said target gene;
 - b)- measuring the transcription or expression of said target gene in the absence and presence of said compound; and
 - c)- comparing the transcription or expression of said target gene in the absence and presence of said compound;

wherein an increase or a decrease in transcription or expression in the presence of said compound indicates that said compound modulates MCHR2.

- 60. (Withdrawn) A method for identifying a compound that selectively modulates MCHR2, comprising
 - a)- contacting an isolated or recombinant MCHR polypeptide with a compound of Claim 1 under conditions suitable for ligand binding to said MCHR;

- 14 -

- b)- measuring the binding affinities of said compound for said MCHR and for an MCHR2 polypeptide; and
- c)- comparing the binding affinities of said compound for said MCHR and for said MCHR2 polypeptide;

wherein a binding affinity for said MCHR2 polypeptide of at least 10-fold greater than the binding affinity for said MCHR indicates that said MCHR2 compound selectively modulates MCHR2.

- 61. (Withdrawn) The method of any one of Claims 57, 58, 59, and 60 wherein said compound that modulates MCHR2 is an MCHR2 antagonist.
- 62. (Withdrawn) A compound identified according to the method of any one of Claims 57, 58, 59 and 60.
- 63. (Withdrawn) A method for identifying a compound that modulates MCHR2, comprising
 - a)- determining the binding mode of a compound of Claim 61 to MCHR2;
 - b)- modifying said compound to provide a test compound that is capable of said binding mode;
 - c)- contacting an isolated or recombinant MCHR2 polypeptide with an MCHR2 ligand in the absence and presence of said test compound under conditions suitable for ligand binding to said polypeptide;
 - d)- measuring ligand binding to said polypeptide in the absence and presence of said test compound; and
 - e)- comparing ligand binding to said polypeptide in the absence and presence of said test compound;

wherein an increase or a decrease in ligand binding in the presence of said test compound indicates that said test compound modulates MCHR2.